WO 2005/069891 PCT/US2005/001393

CLAIMS

WHAT IS CLAIMED IS:

- 1. A method for performing expression proteomic analysis, comprising: inducing a subtle isotope modification in an organism or sample; analyzing said subtle isotope modification with an analytic tool.
- 2. The method of claim 1 wherein said analytic tool is used to monitor turnover of peptides, polypeptides or both.
- 3. The method of claim 1 wherein said analytic tool is configured to calculate an isotope ratio for specific peptides or polypeptides by measuring isotope distributions.
- 4. The method of claim 1 wherein said organism is a mammal.
- 5. The method of claim 1 wherein said organism is a human.
- 6. The method of claim 1 wherein said subtle isotope modification includes swapping of ¹³C for ¹²C, ¹⁸O for ¹⁶O, ¹⁵N for ¹⁴N, deuterium for hydrogen, or combinations thereof.
- 7. The method of claim 1 wherein said subtle isotope ratio modification results in a modification of a ¹³C:¹²C isotope ratio of from about 100:1 to about 200:1.
- 8. The method of claim 1 wherein said subtle isotope modification is induced by a method selected from the group consisting of isotope-coded affinity tag (ICAT), stable isotope labeling with amino acids in cell culture (SILAC), enzymatic exchange, growth in stable isotopes, diet, injection and combinations thereof.
- 9. The method of claim 1 wherein said subtle isotope modification is analyzed by a method selected from the group consisting of matrix-assisted laser desorption ionization time-of-flight (MALDI-TOF) mass spectrometry, micro-liquid chromatography with tandem mass spectrometry (LC-MSMS), high resolution mass spectrometry, tandem mass spectrometry (MSMS), Fournier transform mass spectrometry (FTMS), averagine, isotope ratio mass spectrometry (IRMS) and combinations thereof.

WO 2005/069891 PCT/US2005/001393

- 10. A system for analyzing isotope distribution, comprising:
 an organism in which a subtle isotope modification has been induced; and
 an analytic tool to analyze said subtle isotope modification.
- 11. The system of claim 10 wherein said analytic tool is configured to analyze turnover of peptides, polypeptides or both.
- 12. The system of claim 10 wherein said analytic tool is configured to calculate an isotope ratio for specific peptides or polypeptides by measuring isotope distributions.
- 13. The system of claim 10 wherein said organism is a mammal.
- 14. The system of claim 10 wherein said organism is a human.
- 15. The system of claim 10 wherein said subtle isotope modification includes swapping of ¹³C for ¹²C, ¹⁸O for ¹⁶O, ¹⁵N for ¹⁴N, deuterium for hydrogen, or combinations thereof.
- 16. The system of claim 10 wherein said subtle isotope ratio modification results in a modification of a ¹³C:¹²C isotope ratio of from about 100:1 to about 200:1.
- 17. The system of claim 10 wherein said subtle isotope modification is induced by a method selected from the group consisting of isotope-coded affinity tag (ICAT), stable isotope labeling with amino acids in cell culture (SILAC), enzymatic exchange, growth in stable isotopes, diet, injection and combinations thereof.
- 18. The system of claim 10 wherein said analytic tool is configured to analyze said subtle isotope modification by a method selected from the group consisting of matrix-assisted laser desorption ionization time-of-flight (MALDI-TOF) mass spectrometry, microliquid chromatography with tandem mass spectrometry (LC-MSMS), high resolution mass spectrometry, tandem mass spectrometry (MSMS), Fournier transform mass spectrometry (FTMS), averagine, isotope ratio mass spectrometry (IRMS) and combinations thereof.

WO 2005/069891 PCT/US2005/001393

19. A method for performing expression proteomic analysis, comprising:

providing an elemental composition in which a subtle isotope modification has been induced;

analyzing said subtle isotope modification with the Isosolv algorithm.

- 20. The method of claim 19 wherein said subtle isotope modification includes swapping of ¹³C for ¹²C, ¹⁸O for ¹⁶O, ¹⁵N for ¹⁴N, deuterium for hydrogen, or combinations thereof.
- 21. The method of claim 19 wherein said subtle isotope ratio modification results in a modification of a ¹³C:¹²C isotope ratio of from about 100:1 to about 200:1.
- 22. The method of claim 19 wherein said subtle isotope modification is induced by a method selected from the group consisting of isotope-coded affinity tag (ICAT), stable isotope labeling with amino acids in cell culture (SILAC), enzymatic exchange, growth in stable isotopes, diet, injection and combinations thereof.
- 23. The method of claim 19 wherein said subtle isotope modification is analyzed by a method selected from the group consisting of matrix-assisted laser desorption ionization time-of-flight (MALDI-TOF) mass spectrometry, micro-liquid chromatography with tandem mass spectrometry (LC-MSMS), high resolution mass spectrometry, tandem mass spectrometry (MSMS), Fournier transform mass spectrometry (FTMS), averagine, isotope ratio mass spectrometry (IRMS) and combinations thereof.